PATENT

HORST LINDHOFER et al. Application No.: 09/094,921, Group Art Unit: 1642, Examiner: Holleran, A. Amendment No. 3 -- Page 2

1.	(twice amended) Method for the preparation of a vaccine for immunization of humans
	and animals against tumor cells comprising the steps of:

- a) isolating autologous tumor cells;
- b) treating the tumor cells to prevent the survival thereof following reinfusion;
- c) incubating the thus treated tumor cells with intact heterologous bispecific antibodies showing the following properties:
 - α binding to a T cell;
 - β binding to at least one antigen on a tumor oell;
 - γ binding, by their Fc portion to Fc receptor-positive cells capable of activating the Fc receptor-positive cell whereby the expression of cytokines, co-stimulatory antigens or both is induced or increased, wherein the bispecific antibodies are members selected from the group consisting of the following isotype combinations:

```
rat-IgG2b/human-IgG1/
rat-IgG2b/human-IgG2,
rat-IgG2b/human-IgG3[oriental allotype G3m(st) = binding to protein A],
rat-IgG2b/human-IgG4;
```

rat-IgG2b/rat-IgG2c;

mouse fgG2a/human-IgG3[caucasian allotypes <math>G3m(b+g) = no binding to protein A, in the following indicated as *]

mouse-IgG2a/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3]

mouse-IgG2a/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human- IgG3*-[CH2-CH3]

		HORST LINDHOFER et al. Application No.: 09/094,921, Group Art Unit: 1642, Examiner: Holleran, A. Amendment No. 3 Page 3
	25	mouse-IgG2a/human-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-
	26	[CH2-CH3]
	27	mouse-[VH-CH1,VL-CL]-human-IgG1/rat-[VH-CH1,VL-CL]-
	28	human-IgG1-[hinge]-human-IgG3*-[CH2-CH3]
	29	mouse-[VH-CH1,VL-CL]-human-IgG4/rat-[VH-CH1,VL-CL]-human-IgG4-
	30	[hinge]-human-IgG4[N-terminal region of CH2]-human-IgG3*[C-terminal region
•	31	of CH2: > aa position 251]-human-IgG3*[CH3]
	32	rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge-CH2-CH3]
2	33	rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG2-[hinge-CH2-CH3]
Λ	34	rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG3-[hinge-CH2-CH3, oriental
	35	allotype]
W	36	rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG4-[hinge-CH2-CH3]
	37	human-IgG1/human-[VH-CH1/VL-CL]-human-IgG1-[hinge]-
	38	human-IgG3*-[CH2-CH3]
	39	human-IgG1/rat-[VH-CH],VL-CL]-human-IgG1-[hinge]-human-IgG4[N-
	40	terminal region of CH2] human-IgG3*[C-terminal region of CH2 : > aa position
	41	251]-human-IgG3*[CH3]
	42	human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4[N-
	43	terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position
	44	251]-human-Ig@3*[CH3]
	45	human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG2[N-
	46	terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position
	47	251]-hyman-IgG3*[CH3]
	11	

	HORST LINDHOFER et al. Application No.: 09/094,921, Group Art Unit: 1642, Examiner: Holleran, A. Amendment No. 3 Page 4
48	human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hing#]-human-IgG2[N-
49	terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position
50	251]-human-IgG3*[CH3]
51	human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-
52	CH3]
53	human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-
54	[CH2-CH3]
55	human-IgG2/human-[VH-CH1,VL-CL]-human-IgG2-[hinge]-human-IgG3*-
56	[CH2-CH3]
. 57	human-IgG4/human-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG3*-
∩ / . ⁵⁸	[CH2-CH3]
€ 1 1 59	human-IgG4/human-[VH-CH1, VL-CL]-human-IgG4-[hinge]-human-IgG4[N-
WI 60	terminal region of CH2]-humar-IgG3*[C-terminal region of CH2 : > aa position
61	251]-human-IgG3*[CH3]
62	mouse-IgG2b/rat-[VH-CH/I,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-
63	CH3]
64	mouse-IgG2b/human-fVH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-
65	[CH2-CH3]
. 66	mouse-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-
67	[CH2-CH3]
68	mouse-[VH-CH1,VL-CL]-human-IgG4/rat-[VH-CH1,VL-CL]-human-IgG4-
69	[hinge]-human-IgG4-[CH2]-human-IgG3*-[CH3]
70	human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4-[CH2]-
71	human-IgG3*-[CH3]

, •

7